5

10

15

20

25

30

1/2

Fig 1 A Preparation of inoculum Inoculation of embryonated eggs Incubation Harvest Crude monovalent whole virus bulk Clarification Adsorption Filtration Continuous flow centrifugation Fractionation Monovalent purified whole virus pool Sterile filtration Inactivation Final filtraion

Monovalent final bulk

Storage

Sterile filtration

2/2

Fig 1 B

## Down stream process

Infected allantoic fluid Clarification by centrifugation (500 - 1500 g) Adsorption on CaHPO<sub>4</sub> - gel Elution by EDTA Centrifugation over linear sucrose Collect fractions Sterile filtration Trypsin digestion Inactivation 2 hrs / 37°C 20° ± 2°C, 72 hrs, 250 µg/ml formaldehyde Ultrafiltration Kd = 30 - 50000Inactivation 20° ± 2°C, 72 hrs, 250 µg/ml formaldehyde

Store at 2 - 8°C